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| U.S. DEPARTMENT OF COMMERCE, PATENT AND TRADEMARK OFFICE DATE: March 2, 2006  |  |  |  |  |  |  |  |  |
|---|--|--|--|--|--|--|--|--|
| TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE U.S. Application. Application.                              |  |  |  |  |  |  |  |  |
| (DO/EO/US) CONCERNING A FILING UNDER 35 U.S.C. 371  |  |  |  |  |  |  |  |  |
| INTERNATIONAL APPLICATION NO.: INTERNATIONAL FILING DATE: PRIORITY DATE CLAIMED: March 3, 2004                                |  |  |  |  |  |  |  |  |
| TITLE OF INVENTION: THERAPEUTIC OR PREVENTIVE DRUG FOR OSTEOPOROSIS COMPRISING ISOTAXIRESINOL DERIVED FROM TAXUS YUNNANENSIS  |  |  |  |  |  |  |  |  |
| APPLICANT(S) FOR DO/EO/US: Shigetoshi KADOTA et al.   |  |  |  |  |  |  |  |  |
| Applicant hereby submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: |  |  |  |  |  |  |  |  |
| 1. 🖂  |  |  |  |  |  |  |  |  |
| 2.  | This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371.  |  |  |  |  |  |  |  |
| 3. 🛛  | This is an express request to begin national examination procedures (35 USC 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.   |  |  |  |  |  |  |  |
| 4. 🖾  | The US has been elected (Article 31).  |  |  |  |  |  |  |  |
| 5. 🛛  | A copy of the International Application as filed (35 U.S.C. 371(c)(2)): a. ☐ is transmitted herewith (required only if not transmitted by the International Bureau). b. ☐ has been transmitted by the International Bureau. c. ☐ is not required, as the application was filed in the United States Receiving Office (RO/US)   |  |  |  |  |  |  |  |
| 6. 🛚  | An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).  a. ⊠ is attached hereto.  b. ☐ has been previously submitted under 35 U.S.C. 154(d)(4).  |  |  |  |  |  |  |  |
| 7. 🖾  | Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3)  a. □ are transmitted herewith (required only if not transmitted by the International Bureau).  b. □ have been transmitted by the International Bureau.  c. □ have not been made; however, the time limit for making such amendments has NOT expired.  d. ⊠ have not been made and will not be made. |  |  |  |  |  |  |  |
| 8.  | An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).  |  |  |  |  |  |  |  |
| 9. 🛛  | An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).   |  |  |  |  |  |  |  |
| 10. 🗆   | An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).   |  |  |  |  |  |  |  |
| ITEMS 11. TO 20. BELOW CONCERN OTHER DOCUMENT(S) OR INFORMATION INCLUDED:   |  |  |  |  |  |  |  |  |
| 11. 🛛   | An Information Disclosure Statement under 37 CFR 1.97 and 1.98 together with the international search report, PTO/SB/08 and 4 docs.  |  |  |  |  |  |  |  |
| 12. 🛛   | An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.  ASSIGNEE NAME AND ADDRESS: KOTOSUGI INC., Sagamihara-shi, Japan  Please publish the assignee data with the application.   |  |  |  |  |  |  |  |
| 13. 🗆   | A preliminary amendment.   |  |  |  |  |  |  |  |
| 14. 🛛   | An Application Data Sheet under 37 C.F.R. 1.76   |  |  |  |  |  |  |  |
| 15. 🗆   | A substitute specification.  |  |  |  |  |  |  |  |
| 16. 🗆   | A change of power of attorney and/or address letter.   |  |  |  |  |  |  |  |
| 17. 🗆   | A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 C.F.R. 1.821 - 1.825.  |  |  |  |  |  |  |  |
| 18. 🗆   | A second copy of the published international application under 35 U.S.C. 154(d)(4).  |  |  |  |  |  |  |  |
| 19. 🗆   | A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).  |  |  |  |  |  |  |  |
| 20. 🗆   | Other items or information:  |  |  |  |  |  |  |  |

1AP20 Rec'd FUT/PTO 0 2 MAR 2006
ATTORNEY'S DOCKET NO.: 062043

| U.S. APPLICATION NO. (if known) INTERNATIONAL APPLICATION NO. PCT/JP2005/001055  |  |                                      | DATE: March 2, 2006 | DATE: March 2, 2006 |  |  |  |
|--|--|--------------------------------------|---------------------|---------------------|--|--|--|
| The following fees ar  | re submitted:  | CALCULATIONS                         | PTO USE ONLY        |                     |  |  |  |
|  |  |                                      |                     |                     |  |  |  |
| <b> </b>   | onal Stage Fee   | \$ 300.00                            |                     |                     |  |  |  |
|  | tage Examination Fee   |                                      |                     |                     |  |  |  |
| provisions of PCT Art All other situations   | inary examination repo<br>icle 33(1)-(4)   | \$ 200.00                            |                     |                     |  |  |  |
| Search fee (37 CFR 1.<br>an International Search   | tage Search Fee<br>445(a)(2) has been paid<br>hing Authority<br>eport prepared and pro | O as \$ 400.00                       |                     |                     |  |  |  |
|  | ENT  | ER APPROPRIATE BAS                   | IC FEE AMOU         | VT = \$ 900.00      |  |  |  |
| Surcharge of \$130.00 earliest claimed prior   | o for furnishing the oating the oating the oating the oate (37 CFR 1.49                | the l                                |                     |                     |  |  |  |
| CLAIMS   | NUMBER<br>FILED  | NUMBER EXTRA                         | RATE                |                     |  |  |  |
| TOTAL  | 1 -20 =  | 0                                    | X \$ 50.00          |                     |  |  |  |
| INDEPENDENT  | 1 - 3 =  | 0                                    | X \$ 200.00         |                     |  |  |  |
|  | laims(s) (if applicable  |                                      | + \$360.00          | 6 000 00            |  |  |  |
|  | CALCULATIONS =   | ch additional 50 sheets that         | at exceeds 100 sl   | \$ 900.00           |  |  |  |
| \$ 250.00.   | cation Size rec-lor ca   | cii additional 30 sheets the         | at exceeds 100 si   | icets               |  |  |  |
|  | filing by small entity,  | 8). \$ -450.00                       |                     |                     |  |  |  |
| SUBTOTAL =   |  |                                      |                     | \$ 450.00           |  |  |  |
|  | 10.00 for furnishing the riority date (37 CFR 1  | e English translation later (        | than 30 months      | from                |  |  |  |
| TOTAL NATIONAL   |  | .472(1)). +                          | <del> </del>        | \$ 450.00           |  |  |  |
|  |  | t (37 CFR 1.21(h)). The              | assignment mus      |                     |  |  |  |
| \$40.00 per property   |  | \$ 40.00                             |                     |                     |  |  |  |
| TOTAL FEES ENCL  | OSED =   |                                      | ,                   | \$ 490.00           |  |  |  |
|  |  | Amount to be:<br>refunded<br>charged | \$<br>\$            |                     |  |  |  |
| a. A check in  | the amount of \$490.   | 00 to cover the above fees i         | is enclosed. SM     | ALL ENTITY          |  |  |  |
| (\$150.00 for basic national stage fee; \$200.00 for national stage search fee; \$100.00 for national stage examination fee and \$40.00 for the assignment recordation fee).                                   |  |                                      |                     |                     |  |  |  |
| b. Please charge my Deposit Account No. 50-2866 in the amount of \$ to cover the above fees.   |  |                                      |                     |                     |  |  |  |
| c.  The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 50-2866.   |  |                                      |                     |                     |  |  |  |
| NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed to request that the application be restored to pending status. |  |                                      |                     |                     |  |  |  |
| Send All Correspondence To:  WESTERMAN, HATTORI, DANIELS & ADRIAN, LLP 1250 Connecticut Avenue NW, Suite 700 Washington, DC 20036  |  |                                      |                     |                     |  |  |  |
| Timed to Delic 121   | MENUTE VILLE   | (                                    | CUSTOMER NO.        | 38834               |  |  |  |
| 1 yped or Printed Nai  | me: NEW Ken-Ichi H   | аногі                                | Reg. No.: 32,       | 108                 |  |  |  |
| Signature Date: March 2, 2006  |  |                                      |                     |                     |  |  |  |